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                   USPATFULL/USPAT2
 NEWS  8  MAY 30   The F-Term thesaurus is now available in CA/CAplus
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                   INPADOC
 NEWS 10  JUN 26   TULSA/TULSA2 reloaded and enhanced with new search and
                   and display fields
 NEWS 11  JUN 28   Price changes in full-text patent databases EPFULL and PCTFULL
 NEWS 12  JUl 11   CHEMSAFE reloaded and enhanced
 NEWS 13  JUl 14   FSTA enhanced with Japanese patents
 NEWS 14  JUl 19   Coverage of Research Disclosure reinstated in DWPI
 NEWS 15  AUG 09   INSPEC enhanced with 1898-1968 archive
 NEWS 16  AUG 28   ADISCTI Reloaded and Enhanced
 NEWS 17  AUG 30   CA(SM)/CAplus(SM) Austrian patent law changes

 NEWS EXPRESS  JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT
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=> s wo 0017158/pn
L1            0 WO 0017158/PN
                (WO17158/PN)

=> s wo 20017158/pn
L2            0 WO 20017158/PN
                (WO20017158/PN)

=> s wo 200017158/pn
L3            1 WO 200017158/PN
                (WO2000017158/PN)

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L3   ANSWER 1 OF 1  CAPLUS  COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:         2000:210111  CAPLUS
DOCUMENT NUMBER:          132:222873
ENTRY DATE:               Entered STN:  31 Mar 2000
TITLE:                    Preparation of 3-amidinophenylalanine peptides for use
                          as urokinase inhibitors
INVENTOR(S):              Wikstrom, Peter; Vieweg, Helmut
PATENT ASSIGNEE(S):       Pentapharm A.-G., Switz.
SOURCE:                   PCT Int. Appl., 30 pp.
                          CODEN: PIXXD2
DOCUMENT TYPE:            Patent
LANGUAGE:                 German
INT. PATENT CLASSIF.:
            MAIN:         C07C311-19
       SECONDARY:         C07D295-20; C07D211-60; C07D211-62; C07D211-16;
                          C07D409-12; C07D295-18; A61K031-445; A61K031-50
CLASSIFICATION:           34-3 (Amino Acids, Peptides, and Proteins)
                          Section cross-reference(s): 1
FAMILY ACC. NUM. COUNT:   1
PATENT INFORMATION:

     PATENT NO.           KIND   DATE        APPLICATION NO.         DATE
     ---------------      ----   --------    --------------------    --------
     WO 2000017158         A1    20000330    WO 1998-CH402           19980918 <--
         W:  AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG,
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KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

AU 9890597	A1	20000410	AU 1998-90597		19980918
EP 1114024	A1	20010711	EP 1998-942443		19980918
EP 1114024	B1	20021127			
R: DE, ES, FR, GB, IT					
ES 2188003	T3	20030616	ES 1998-942443		19980918
PRIORITY APPLN. INFO.:			EP 1998-942443	A	19980918
			WO 1998-CH402	A	19980918

PATENT CLASSIFICATION CODES:

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2000017158	ICM	C07C311-19
	ICS	C07D295-20; C07D211-60; C07D211-62; C07D211-16; C07D409-12; C07D295-18; A61K031-445; A61K031-50
	IPCI	C07C0311-19 [ICM,6]; C07C0311-00 [ICM,6,C*]; C07D0295-20 [ICS,6]; C07D0211-60 [ICS,6]; C07D0211-62 [ICS,6]; C07D0211-16 [ICS,6]; C07D0211-00 [ICS,6,C*]; C07D0409-12 [ICS,6]; C07D0409-00 [ICS,6,C*]; C07D0295-18 [ICS,6]; C07D0295-00 [ICS,6,C*]; A61K0031-445 [ICS,6]; A61K0031-50 [ICS,6]
	IPCR	C07C0311-00 [I,C*]; C07C0311-19 [I,A]; C07D0211-00 [I,C*]; C07D0211-16 [I,A]; C07D0211-60 [I,A]; C07D0211-62 [I,A]; C07D0295-00 [I,C*]; C07D0295-185 [I,A]; C07D0295-205 [I,A]; C07D0409-00 [I,C*]; C07D0409-12 [I,A]
	ECLA	C07C311/19; C07D211/16; C07D211/60; C07D211/62; C07D295/18B1G; C07D295/20B1; C07D409/12+333B+211
AU 9890597	IPCI	C07C0311-19 [ICM,6]; C07C0311-00 [ICM,6,C*]; C07D0295-20 [ICS,6]; C07D0211-60 [ICS,6]; C07D0211-62 [ICS,6]; C07D0211-16 [ICS,6]; C07D0211-00 [ICS,6,C*]; C07D0409-12 [ICS,6]; C07D0409-00 [ICS,6,C*]; C07D0295-18 [ICS,6]; C07D0295-00 [ICS,6,C*]; A61K0031-445 [ICS,6]; A61K0031-50 [ICS,6]
	IPCR	C07C0311-00 [I,C*]; C07C0311-19 [I,A]; C07D0211-00 [I,C*]; C07D0211-16 [I,A]; C07D0211-60 [I,A]; C07D0211-62 [I,A]; C07D0295-00 [I,C*]; C07D0295-185 [I,A]; C07D0295-205 [I,A]; C07D0409-00 [I,C*]; C07D0409-12 [I,A]
EP 1114024	IPCI	C07C0311-19 [ICM,6]; C07C0311-00 [ICM,6,C*]; C07D0295-20 [ICS,6]; C07D0211-60 [ICS,6]; C07D0211-62 [ICS,6]; C07D0211-16 [ICS,6]; C07D0211-00 [ICS,6,C*]; C07D0409-12 [ICS,6]; C07D0409-00 [ICS,6,C*]; C07D0295-18 [ICS,6]; C07D0295-00 [ICS,6,C*]; A61K0031-445 [ICS,6]; A61K0031-50 [ICS,6]
	IPCR	C07C0311-00 [I,C*]; C07C0311-19 [I,A]; C07D0211-00 [I,C*]; C07D0211-16 [I,A]; C07D0211-60 [I,A]; C07D0211-62 [I,A]; C07D0295-00 [I,C*]; C07D0295-185 [I,A]; C07D0295-205 [I,A]; C07D0409-00 [I,C*]; C07D0409-12 [I,A]
ES 2188003	IPCI	C07C0311-19 [ICM,4]; C07C0311-00 [ICM,4,C*]; C07D0295-20 [ICS,4]; C07D0211-60 [ICS,4]; C07D0211-62 [ICS,4]; C07D0211-16 [ICS,4]; C07D0211-00 [ICS,4,C*]; C07D0409-12 [ICS,4]; C07D0409-00 [ICS,4,C*]; C07D0295-18 [ICS,4]; C07D0295-00 [ICS,4,C*]; A61K0031-445 [ICS,4]; A61K0031-50 [ICS,4]
	IPCR	A61K0031-445 [I,A]; A61K0031-445 [I,C*]; A61K0031-50 [I,A]; A61K0031-50 [I,C*]; C07C0311-00 [I,C*]; C07C0311-19 [I,A]; C07D0211-00 [I,C*]; C07D0211-16 [I,A]; C07D0211-60 [I,A]; C07D0211-62 [I,A];

C07D0295-00 [I,C*]; C07D0295-18 [I,A]; C07D0295-20 [I,A]; C07D0409-00 [I,C*]; C07D0409-12 [I,A]

OTHER SOURCE(S): MARPAT 132:222873

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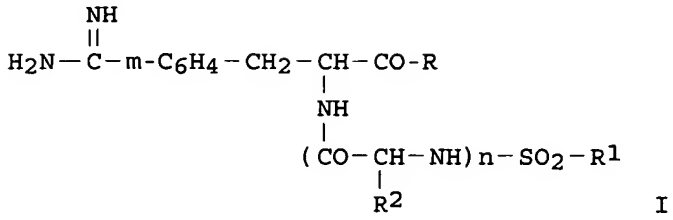

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ABSTRACT:

Title compds. [(I); R = OH, O-(cyclo)alkyl, O-arylalkyl, PhCH2, Ph(CH2)2, substituted pyrrolidine, piperidine, piperazine, NR3R4; R3, R4 = (independently) H, (un)branched alkyl, (un)substituted aralkyl, PhCH2, Ph(CH2)2, cycloalkyl-alkyl; R3 = H, R4 = NHR5; R5 = (hetero)aryl; R1 = (un)branched alkyl, (un)substituted (hetero)aryl; R2 = H, (un)branched alkyl; n = 0-1] as L-,D-, or DL forms, were prepared for use as urokinase inhibitors for the treatment of tumors or in diagnosis. Thus, (L)-3-cyanophenylalanine Me ester hydrochloride was N-protected with 2,4,6-triisopropylphenylsulfonyl chloride, deesterified, condensed with 1-ethoxycarbonyl-piperazine, and the cyano group converted to the amidine (via conversion to thioamide and reaction with MeI to give thioimide Me ester, which was then reacted with ammonium acetate), to give I [R = 4-ethoxycarbonyl-piperazine; R1 = 2,4,6-triisopropylphenyl; n = 0 (II)]. In in vivo tests of urokinase inhibition, II had Ki 0.49 μmol/l.

SUPPL. TERM: amidinophenylalanine peptide prepn urokinase inhibitor tumor

INDEX TERM: Enzyme kinetics
(of inhibition; preparation of amidinophenylalanine peptides for use as urokinase inhibitors)

INDEX TERM: Neoplasm
(preparation of amidinophenylalanine peptides for use as urokinase inhibitors)

INDEX TERM: Peptides, preparation
ROLE: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of amidinophenylalanine peptides for use as urokinase inhibitors)

INDEX TERM: 161357-71-7P 169388-44-7P 220355-61-3P 220355-63-5P
220355-64-6P 222842-26-4P 255374-84-6P 255374-89-1P
255374-90-4P 256430-86-1P 256430-96-3P 261158-90-1P
261158-91-2P 261158-92-3P 261158-93-4P 261158-94-5P
261158-95-6P 261158-96-7P
ROLE: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of amidinophenylalanine peptides for use as urokinase inhibitors)

INDEX TERM: 120-43-4 5006-62-2 6553-96-4 161357-88-6 177740-32-8
ROLE: RCT (Reactant); RACT (Reactant or reagent)
(preparation of amidinophenylalanine peptides for use as urokinase inhibitors)

INDEX TERM: 255374-79-9P 255374-80-2P 255374-81-3P 255374-82-4P
255374-83-5P 255374-85-7P 255374-86-8P 255374-87-9P

255374-88-0P 255375-04-3P
ROLE: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of amidinophenylalanine peptides for use as urokinase inhibitors)

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD.

REFERENCE(S):

(1) Markwardt, F; THROMBOSIS RESEARCH 1980, V17(3), P425
(2) Markwardt, F; THROMBOSIS RESEARCH 1980, V17(3), P425
(3) StUrzebecher, J; BIOLOGICAL CHEMISTRY HOPPE-SEYLER 1997, V373(10), P1025
(4) StUrzebecher, J; BIOLOGICAL CHEMISTRY HOPPE-SEYLER 1997, V373(10), P1025
(5) StUrzebecher, J; JOURNAL OF MEDICINAL CHEMISTRY 1997, V19(40), P3091
(6) StUrzebecher, J; JOURNAL OF MEDICINAL CHEMISTRY 1997, V19(40), P3091
(7) StUrzenbecher, J; DIE PHARMAZIE 1981, V36(7), P501
(8) StUrzenbecher, J; DIE PHARMAZIE 1981, V36(7), P501
(9) Wagner, G; DIE PHARMAZIE 1981, V36(7), P467 CAPLUS
(10) Wagner, G; DIE PHARMAZIE 1981, V36(7), P467 CAPLUS

=> s wo 200004954/pn
L4 2 WO 200004954/PN
(WO2000004954/PN)

=> d l4 iall 1-2

L4 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2004:569861 CAPLUS
DOCUMENT NUMBER: 141:99681
ENTRY DATE: Entered STN: 16 Jul 2004
TITLE: 3-Amidinophenylalanine derivatives as urokinase inhibitors for treatment of cancer
INVENTOR(S): Wilhelm, Olaf; Magdolen, Viktor; Sturzebecker, Jorg; Foekens, John; Lutz, Verena
PATENT ASSIGNEE(S): Wilex A.-G., Germany
SOURCE: U.S. Pat. Appl. Publ., 36 pp., Cont.-in-part of U.S. Ser. No. 202,850.
CODEN: USXXCO
DOCUMENT TYPE: Patent
LANGUAGE: English
INT. PATENT CLASSIF.:
MAIN: A61K031-495
SECONDARY: A61K031-704; A61K031-337
US PATENT CLASSIF.: 514255010; 514034000; 514050000; 514449000; 514414000; 424649000
CLASSIFICATION: 1-6 (Pharmacology)
Section cross-reference(s): 34
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004138233	A1	20040715	US 2003-691528	20031024
WO 2000004954	A2	20000203	WO 1999-EP5145	19990720 <--
WO 2000004954	A3	20000622		

W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,

ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

EP 1264828	A1	20021211	EP 2002-16038	19990720
EP 1264828	B1	20040414		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY				
US 6624169	B1	20030923	US 2001-743800	20010403
US 2003013723	A1	20030116	US 2002-202850	20020726
US 6680320	B2	20040120		

PRIORITY APPLN. INFO.:

EP 1998-113519	A 19980720
WO 1999-EP5145	W 19990720
US 2001-743800	A3 20010403
US 2002-202850	A2 20020726
EP 1999-936570	A3 19990720

PATENT CLASSIFICATION CODES:

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
US 2004138233	ICM	A61K031-495
	ICS	A61K031-704; A61K031-337
	INCL	514255010; 514034000; 514050000; 514449000; 514414000; 424649000
	IPCI	A61K0031-495 [ICM,7]; A61K0031-704 [ICS,7]; A61K0031-7028 [ICS,7,C*]; A61K0031-337 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	NCL	514/255.010; 424/649.000; 514/034.000; 514/050.000; 514/414.000; 514/449.000
	ECLA	A61K031/495+A; A61K051/04; C07D295/20B1
WO 2000004954	IPCI	A61R0031-495 [ICM,6]; A61R0031-445 [ICS,6]; A61R0031-195 [ICS,6]; C07D0295-182 [ICS,6]; C07D0295-00 [ICS,6,C*]; A61R0047-48 [ICS,6]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; C07D295/20B1; A61K051/04G
EP 1264828	IPCI	C07D0295-18 [ICM,6]; C07D0295-00 [ICM,6,C*]; A61K0031-495 [ICS,6]; A61P0035-00 [ICS,6]
	ECLA	C07D295/20B1
US 6624169	IPCI	A61K0031-495 [ICM,7]; A61K0031-445 [ICS,7]; A61K0031-195 [ICS,7]; A61K0031-185 [ICS,7,C*]; A61K0047-48 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	NCL	514/255.010; 514/330.000; 544/388.000; 546/226.000
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
US 2003013723	IPCI	A61K0031-495 [ICM,7]; C07D0241-04 [ICS,7]; C07D0241-00 [ICS,7,C*]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	NCL	514/255.010; 544/386.000
	ECLA	A61K031/495+A; A61K051/04; C07D295/20B1

OTHER SOURCE(S): MARPAT 141:99681

ABSTRACT:
The invention discloses the use of 3-amidinophenylalanine derivs. as urokinase inhibitors for treating malignant tumors and the formation of metastases. Preparation of Nα-2,4,6-triisopropylphenylsulfonyl- DL-3-cyanophenylalanylnipecotic acid benzylamide is described.

SUPPL. TERM: urokinase inhibitor prepn amidinophenylalanine deriv cancer metastasis

INDEX TERM: Drug interactions

(additive; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Lymph node
(axillary, i.p.; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Mammary gland, neoplasm
(carcinoma; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Drug delivery systems
(carriers; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Carcinoma
(mammary; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Fibrins
ROLE: BSU (Biological study, unclassified); BIOL (Biological study)
(matrix degradation by breast carcinoma cells; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Neoplasm
(metastasis; preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Antitumor agents
Cytotoxic agents
Drug bioavailability
Drug delivery systems
Human
Lung, neoplasm
Lymph node
Lymphatic system
Mammary gland, neoplasm
Neoplasm
Pancreas, neoplasm
Radiotherapy
Surgery
(preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: Isotopomers
Taxanes
ROLE: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 220355-63-5P
ROLE: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 255374-84-6P 255374-90-4P
ROLE: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 51-21-8 15663-27-1, Cisplatin 23214-92-8, Doxorubicin

33069-62-4, Paclitaxel 41575-94-4, Carboplatin
56420-45-2, Epirubicin
ROLE: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 74-88-4, Methyl iodide, reactions 120-43-4, 1-Ethoxycarbonylpiperazine 5006-62-2, Ethyl nipecotate 6553-96-4 255374-78-8
ROLE: RCT (Reactant); RACT (Reactant or reagent)
(3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 255374-79-9P 255374-80-2P 255374-81-3P 255374-85-7P 255374-86-8P 255375-04-3P
ROLE: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 255374-89-1P
ROLE: SPN (Synthetic preparation); PREP (Preparation)
(3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 9039-53-6, Urokinase
ROLE: BSU (Biological study, unclassified); BIOL (Biological study)
(inhibitor; 3-amidinophenylalanine derivs. preparation as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 220355-63-5D, mixture with taxanes 255374-90-4D, mixture with taxanes 721396-20-9 721396-21-0 721396-22-1 721396-23-2 721396-24-3 721396-25-4 721396-26-5 721396-27-6 721396-28-7 721396-29-8 721396-30-1 721396-31-2
ROLE: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

INDEX TERM: 255374-87-9P
ROLE: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of 3-amidinophenylalanine derivs. as urokinase inhibitors for treatment of cancer and metastases)

L4 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2000:84664 CAPLUS
DOCUMENT NUMBER: 132:108297
ENTRY DATE: Entered STN: 04 Feb 2000
TITLE: Preparation and use of urokinase inhibitors in the treatment of malignant tumors
INVENTOR(S): Wilhelm, Olaf; Magdolen, Viktor; Sturzebecher, Jorg; Foekens, John; Lutz, Verena
PATENT ASSIGNEE(S): Wilex Biotechnology Gmbh, Germany
SOURCE: PCT Int. Appl., 41 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: German
INT. PATENT CLASSIF.:
MAIN: A61R031-495
SECONDARY: A61R031-445; A61R031-195; C07D295-182; A61R047-48
CLASSIFICATION: 34-2 (Amino Acids, Peptides, and Proteins)
Section cross-reference(s): 1, 63
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000004954	A2	20000203	WO 1999-EP5145	19990720 <--
WO 2000004954	A3	20000622		
W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
CA 2338073	AA	20000203	CA 1999-2338073	19990720
AU 9951615	A1	20000214	AU 1999-51615	19990720
AU 754958	B2	20021128		
BR 9912327	A	20010502	BR 1999-12327	19990720
EP 1098651	A2	20010516	EP 1999-936570	19990720
EP 1098651	B1	20030108		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
JP 2002521348	T2	20020716	JP 2000-560945	19990720
EP 1264828	A1	20021211	EP 2002-16038	19990720
EP 1264828	B1	20040414		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, CY				
AT 230599	E	20030115	AT 1999-936570	19990720
ES 2189455	T3	20030701	ES 1999-936570	19990720
AT 264103	E	20040415	AT 2002-16038	19990720
PT 1264828	T	20040831	PT 2002-16038	19990720
ES 2219607	T3	20041201	ES 2002-16038	19990720
US 6624169	B1	20030923	US 2001-743800	20010403
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US 6680320	B2	20040120		
US 2004138233	A1	20040715	US 2003-691528	20031024
PRIORITY APPLN. INFO.:			EP 1998-113519 A	19980720
			EP 1999-936570 A3	19990720
			WO 1999-EP5145 W	19990720
			US 2001-743800 A3	20010403
			US 2002-202850 A2	20020726

PATENT CLASSIFICATION CODES:

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2000004954	ICM	A61R031-495
	ICS	A61R031-445; A61R031-195; C07D295-182; A61R047-48
	IPCI	A61R0031-495 [ICM,6]; A61R0031-445 [ICS,6]; A61R0031-195 [ICS,6]; C07D0295-182 [ICS,6]; C07D0295-00 [ICS,6,C*]; A61R0047-48 [ICS,6]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; C07D295/20B1; A61K051/04G
CA 2338073	IPCI	A61K0031-495 [ICM,6]; C07D0295-182 [ICS,6]; C07D0295-00 [ICS,6,C*]; A61K0031-195 [ICS,6]; A61K0031-185 [ICS,6,C*]; A61K0031-445 [ICS,6]; A61K0047-48 [ICS,6]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
AU 9951615	IPCI	C07D0295-182 [ICM,6]; C07D0295-00 [ICM,6,C*]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
BR 9912327	IPCI	A61K0031-495 [ICM,7]; A61K0031-445 [ICS,7];

		A61K0031-195 [ICS,7]; A61K0031-185 [ICS,7,C*]; C07D0295-182 [ICS,7]; C07D0295-00 [ICS,7,C*]; A61K0047-48 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
EP 1098651	IPCI	A61K0031-495 [ICM,6]
	IPCR	C07D0295-00 [I,C*]; C07D0295-205 [I,A]
JP 2002521348	IPCI	A61K0031-445 [ICM,7]; A61K0009-02 [ICS,7]; A61K0009-06 [ICS,7]; A61K0009-08 [ICS,7]; A61K0009-14 [ICS,7]; A61K0009-20 [ICS,7]; A61K0009-48 [ICS,7]; A61K0031-495 [ICS,7]; A61P0017-00 [ICS,7]; A61P0035-00 [ICS,7]; A61P0043-00 [ICS,7]; C07D0211-60 [ICS,7]; C07D0211-00 [ICS,7,C*]; C07D0295-20 [ICS,7]; C07D0295-00 [ICS,7,C*]; C07M0007-00 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
EP 1264828	IPCI	C07D0295-18 [ICM,6]; C07D0295-00 [ICM,6,C*]; A61K0031-495 [ICS,6]; A61P0035-00 [ICS,6]
	ECLA	C07D295/20B1
AT 230599	IPCI	A61K0031-495 [ICM,7]; A61P0035-00 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
ES 2189455	IPCI	A61K0031-495 [ICM,4]; A61P0035-00 [ICS,4]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
AT 264103	IPCI	A61K0031-495 [ICM,7]; A61P0035-00 [ICS,7]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
PT 1264828	IPCI	A61K0031-495 [ICM,7]; A61P0035-00 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
ES 2219607	IPCI	A61K0031-495 [ICM,7]; A61P0035-00 [ICS,4]
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
US 6624169	IPCI	A61K0031-495 [ICM,7]; A61K0031-445 [ICS,7]; A61K0031-195 [ICS,7]; A61K0031-185 [ICS,7,C*]; A61K0047-48 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	NCL	514/255.010; 514/330.000; 544/388.000; 546/226.000
	ECLA	A61K031/495+A; A61K051/04; A61K051/04G; C07D295/20B1
US 2003013723	IPCI	A61K0031-495 [ICM,7]; C07D0241-04 [ICS,7]; C07D0241-00 [ICS,7,C*]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	NCL	514/255.010; 544/386.000
	ECLA	A61K031/495+A; A61K051/04; C07D295/20B1
US 2004138233	IPCI	A61K0031-495 [ICM,7]; A61K0031-704 [ICS,7]; A61K0031-7028 [ICS,7,C*]; A61K0031-337 [ICS,7]
	IPCR	A61K0031-495 [I,A]; A61K0031-495 [I,C*]; A61K0051-02 [I,C*]; A61K0051-04 [I,A]; C07D0295-00 [I,C*]; C07D0295-205 [I,A]
	NCL	514/255.010; 424/649.000; 514/034.000; 514/050.000; 514/414.000; 514/449.000
	ECLA	A61K031/495+A; A61K051/04; C07D295/20B1

OTHER SOURCE(S): MARPAT 132:108297

ABSTRACT:
The invention relates to the use of derivs. of 3-amidino-phenyl-alanine [H2NC(:NH)-3-C6H4-CH2CH(NH(COCH(R2)NH)nSO2R1)COR; R = OH, (substituted) ester, (substituted) amine, (substituted) heterocycle; R1 = substituted phenyl; R2 = H, (un)branched alkyl; n=0,1 (I)] as urokinase inhibitors for treating malignant tumors and the formation of metastases thereof. Thus, beginning with (L)-3-cyanophenylalanine Me ester and 2,4,6-tri(isopropyl)benzenesulfonyl chloride, (S)-I [n=0; R = 4-ethoxycarbonyl-piperazinyl; R1 = 2,4,6-tri(isopropyl)-C6H2; (II)] was synthesized in four steps. In in vitro inhibition tests of urokinase, II had Ki 0.41 μM/l; the compound prepared from (DL)-phenylalanine starting material had Ki 0.96 μM/l.

SUPPL. TERM: amidinophenylalanine deriv urokinase inhibitor prepn cancer tumor metastasis treatment

INDEX TERM: Neoplasm
(metastasis; preparation and use of amidinophenylalanine derivs. as urokinase inhibitors in the treatment of malignant tumors to limit metastasis)

INDEX TERM: Mammary gland
(neoplasm; preparation and use of amidinophenylalanine derivs. as urokinase inhibitors in the treatment of malignant tumors)

INDEX TERM: Skin, disease
(pemphigus vulgaris; preparation and use of amidinophenylalanine derivs. as urokinase inhibitors in the treatment of malignant tumors)

INDEX TERM: Enzyme kinetics
Neoplasm
Pancreas, neoplasm
(preparation and use of amidinophenylalanine derivs. as urokinase inhibitors in the treatment of malignant tumors)

INDEX TERM: Amino acids, preparation
ROLE: RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
(preparation and use of amidinophenylalanine derivs. as urokinase inhibitors in the treatment of malignant tumors)

INDEX TERM: 9039-53-6
ROLE: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
(inhibition of by amidinophenylalanine derivs. for use in the treatment of malignant tumors)

INDEX TERM: 255374-84-6P
ROLE: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation and reaction of in the synthesis of amidinophenylalanine derivs. for use as urokinase inhibitors)

INDEX TERM: 255374-79-9P 255374-80-2P 255374-81-3P 255374-82-4P
255374-83-5P 255374-85-7P 255374-86-8P 255374-87-9P
255374-88-0P 255375-04-3P
ROLE: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation and reaction of in the synthesis of amidinophenylalanine derivs. for use as urokinase inhibitors)

INDEX TERM: 255374-89-1P
ROLE: BAC (Biological activity or effector, except adverse);

BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of for use as urokinase inhibitors)

INDEX TERM: 220355-63-5 255374-90-4
ROLE: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(preparation of for use as urokinase inhibitors)

INDEX TERM: 120-43-4 5006-62-2, Nipecotic acid, ethyl ester
6553-96-4 161357-88-6 255374-78-8
ROLE: RCT (Reactant); RACT (Reactant or reagent)
(reaction of in the synthesis of amidinophenylalanine derivs. for use as urokinase inhibitors)